Applicant or Patentee:

Phillip Dan Cook, Daniel Peter Claude McGee and Charles Guinosso

Serial or Patent No.: N/A

Attorney's Docket No.: ISIS-0710

Date Filed or Issued: HEREWITH

For: NOVEL 2' 0-ALKYL GUANOSINES

VERIFIED STATEMENT (DECLARATION) CLAIMING SMALL ENTITY STATUS
(37 CFR 1.9(f) and 1.27(c) - SMALL BUSINESS CONCERN

I hereby declare that I am:

- () the owner of the small business concern identified below:
- (XX) an official empowered to act on behalf of the concern identified below:

NAME OF CONCERN: Isis Pharmaceuticals ADDRESS OF CONCERN: 2280 Faraday Avenue Carlsbad, CA 92008

I hereby declare that the above-identified small business concern qualifies as a small business concern as defined in 13 CFR 121.12, and reproduced in 37 CFR 1.9(d), for purposes of paying reduced fees under section 41(a) and (b) of Title 35, United States Code, in that: (1) the number of employees of the concern, including those of its affiliates, does not exceed 500 persons; and (2) the concern has not assigned, granted, conveyed, or licensed, and is under no obligation under contract or law to assign, grant, convey, or license, any rights in the invention to any person who could not be classified as an independent inventor if that person had made the invention, or to any concern which would not qualify as a small business concern or a nonprofit organization under this section. For purposes of this statement, (1) the number of employees of the business concern is the average over the previous fiscal year of the concern of the persons employed on a full-time, part-time or temporary basis during each of the pay periods of the fiscal year, and (2) concerns are affiliates of each other when either, directly or indirectly, one concern controls or has the power to control the other, or a third party or parties controls or has the power to control both.

I hereby declare that rights under contract or law have been conveyed to and remain with the small business concern identified above with regard to the invention entitled NOVEL 2' 0-ALKYL GUANOSINES by inventor(s) Phillip Dan Cook, Daniel Peter Claude McGee and Charles Guinosso described in

(XX)	specification	filed	herewith.	

() application	serial no.	, filed	
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() patent no, issued

If the rights held by the above-identified small business concern are not exclusive, each individual, concern or organization having rights to the invention is listed below* and no rights to the invention are held by any person, other than the inventor, who could not qualify as a small business concern under 37 CFR 1.9(d) or by any concern which would not qualify as a small business concern under 37 CFR 1.9(d) or a nonprofit organization under 37 CFR 1.9(e).

*NOTE: Separate verified statements are required for each named person, concern or organization having rights to the invention averring to their status as small entities. (37 CFR 1.27)

FULL NAME:

ADDRESS:

() INDIVIDUAL () SMALL BUSINESS CONCERN () NONPROFIT ORGANIZATION

FULL NAME:

ADDRESS:

() INDIVIDUAL () SMALL BUSINESS CONCERN () NONPROFIT ORGANIZATION

I acknowledge the duty to file, in this application or patent, notification of any change in status resulting in loss of entitlement to small entity status prior to paying, or at the time of paying, the earliest of the issue fee or any maintenance fee due after the date on which status as a small entity is no longer appropriate. (37 CFR 1.28(b))

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application, any patent issuing thereon, or any patent to which this verified statement is directed.

NAME OF PERSON SIGNING B. Lynne Parshall
TITLE OF PERSON SIGNING Vice President and General Counsel
ADDRESS OF PERSON SIGNING Isis Pharmaceuticals, Inc.
2280 Faraday Avenue, Carlsbad, CA 92008

SIGNATURE 2200

Oct 26, 1892

DATE

ROOM POCT PROCKET

NO.: ISIS-0710

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re patent application of:

Cook et al.

Serial No.: N/A Group No.: N/A

Filed: October 27, 1992 Examiner: N/A

For: Novel 2' O-Alkyl Guanosines

Commissioner of Patents & Trademarks Washington, DC 20231

Sir:

ASSOCIATE POWER OF ATTORNEY

The undersigned, Rebecca R. Gaumond, Registration No. 35,152, of the firm WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS, One Liberty Place - 46th Floor, Philadelphia, Pennsylvania 19103, Attorney and/or Agent for Applicant(s), hereby appoints the following:

Robert B. Washburn Richard E. Kurtz John J. Mackiewicz Norman L. Norris Albert W. Preston, Jr. Dale M. Heist Philip S. Johnson John W. Caldwell Gary H. Levin Steven J. Rocci Dianne B. Elderkin Francis A. Paintin John P. Donohue, Jr. Henrik D. Parker Liza D. Hohenschutz	Registration No. 16,574 Registration No. 19,263 Registration No. 19,709 Registration No. 24,196 Registration No. 25,366 Registration No. 28,425 Registration No. 27,200 Registration No. 28,937 Registration No. 28,734 Registration No. 30,489 Registration No. 19,386 Registration No. 29,916 Registration No. 31,863 Registration No. 33,712
	
Dianne B. Elderkin	Registration No. 28,598,
Francis A. Paintin	Registration No. 19,386
John P. Donohue, Jr.	Registration No. 29,916
	Registration No. 31,863,
	Registration No. 33,712
Suzanne E. Miller	Registration No. 32.279
Albert T. Keyack	Registration No. 32,906
Lynn B. Morreale	Registration No. <u>32,842</u>
Joseph Lucci	Registration No. 33,307
Michael P. Dunnam	Registration No. 32,611
Michael D. Stein	Registration No. 34,734
Albert J. Marcellino	Registration No. 34,664

David R. Bailey Registration No. 35,057
Jane Massey Licata Registration No. 32,257
John L. Knoble Registration No. 32,387
Mark DeLuca Registration No. 33,229
Joanne Longo Feeney Registration No. 35,134
Lori Y. Beardell Registration No. 34,293

her associates with full power to prosecute the above-identified application and to transact all business in the Patent Office connected therewith and requests that correspondence continue to be directed to the firm of WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS at the above address.

Date: October 27, 1992

Rebecca R. Gaumond
Attorney of Record

Registration No. 35,152

WOODCOCK WASHBURN KURTZ
MACKIEWICZ & NORRIS
One Liberty Place - 46th Floor
Philadelphia, PA 19103
(215) 568-3100

DOCKET NO.: ISIS-0710 PATENT

COMBINED DECLARATION AND POWER OF ATTORNEY

As a below named inventor, I hereby declare that: My residence, post office address and citizenship are as stated below next to my name; and I verily believe that I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled: NOVEL 2' 0-ALKYL GUANOSINES specification of which: (xx) is attached hereto. () was filed on _____ as Application Serial No. _ and was amended on _____ (if applicable). I hereby state that I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above. I acknowledge the duty to disclose to the U.S. Patent and Trademark Office all information known to be material to the patentability of this application in accordance with 37 CFR § 1.56. I hereby claim foreign priority benefits under 35 U.S.C. § 119 of any foreign application(s) for patent or inventor's certificate listed below and have also identified below any foreign application for patent or inventor's certificate having a filing date before that of any application on which priority is claimed: Country Number Date Filed Priority Claimed

I hereby claim the benefit under 35 U.S.C. § 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of 35 U.S.C. § 112, I acknowledge the duty to disclose to the U.S. Patent and Trademark Office all information known to be material to patentability as defined in 37 CFR § 1.56 which became available between the filing date of the prior application and the national or PCT international filing date of this application:

Application Serial No.	Filing Date	Status (patented, pending)
918,362	July 23, 1992	pending

I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith: John W. Caldwell and Rebecca R. Gaumond, Registration Nos. 28,937 and 35,152 of the firm of WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS, One Liberty Place - 46th Floor, Philadelphia, Pennsylvania 19103, and Herb Boswell, Registration No. 27,311, of ISIS Pharmaceuticals, 2280 Faraday Avenue, Carlsbad, California 92008.

Address all telephone calls and correspondence to:

Rebecca R. Gaumond, Esq.

WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS

One Liberty Place - 46th Floor

Philadelphia, PA 19103

Telephone No. 215-568-3100.

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the

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United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

_00		Full Name Phillip Dan Cook	Anventor's Signature	Date 10/23/92
	1	Residence 7340 Bolero Street Carlsbad, CA 92009		us /
		Post Office Address Same as above		
-00		Full Name <u>Daniel Peter Claude McGee</u>	Inventor's Signature	Date 23 Oct 1992
	2	Residence 295 Chinquapin Aver Carlsbad, CA 92008		nadian
Post Office Address Same as above				
320		Full Name Charles John Guinosso	Inventor's Signature	Date 23001.92
	3	Residence 3 399 Calle de Sur ³ Ca rlsbad, CA 9200	Osia, ca.	us
	Post Office Address Same as above			
		Full Name	Inventor's Signature	Date
	4	Residence	Citizenship	
		Post Office Address		
		Full Name	Inventor's Signature	Date
	Residence Citizenship			
		Post Office Address		